



# Tuberculosis

# Tuberculosis

## ❖ Definition :-

Tuberculosis is a communicable, chronic granulomatous disease, caused by mycobacterium tuberculosis. It usually involves the lungs but may affect any organ or tissue of the body.

## ➤ Etiology:- three types of mycobacterium;

- Mycobacterium tuberculosis: - is the most common cause of infection in human.
- Tuberculin boves :- (reservoir cattle) may the cause of infection in human who use unpasteurized milk.
- *M. africanum* (reservoir humans).
- A typical or opportunistic mycobacterium affecting immuno-compromised human.

# Tuberculosis

## ❖ Pathology and pathogenesis

- *M. bovis* infection arises mainly drinking non-sterilized milk from infected COWS.
- *M. tuberculosis* is spread by the inhalation of aerosolized droplet nuclei from other infected patients.
- Once inhaled, the organisms lodge in the alveoli and initiate the recruitment of macrophages and lymphocytes.
- Macrophages undergo transformation into epithelioid and Langhans cells, which aggregate with the lymphocytes to form the classical tuberculous granuloma.

# Tuberculosis

## ❖ *Pathology and pathogenesis*

- Numerous granulomas aggregate to form a primary lesion or 'Ghon focus' (a pale yellow, caseous nodule, usually a few millimeters to 1–2 cm in diameter), which is characteristically situated in the periphery of the lung.
- Spread of organisms to the hilar lymph nodes is followed by a similar pathological reaction, and the combination of the primary lesion and regional lymph nodes is referred to as the 'primary complex of Ranke'.
- Reparative processes encase the primary complex in a fibrous capsule, limiting the spread of bacilli.
- If no further complications ensue, this lesion eventually calcifies and is clearly seen on a chest X-ray.

# Tuberculosis

## ❖ *Pathology and pathogenesis:-*

- Lymphatic or hematogenous spread may occur before immunity is established, however, seeding secondary foci in other organs, which may lie dormant for years.
- The only clue that infection has occurred may be the appearance of a cell-mediated, delayed-type hypersensitivity reaction to tuberculin, demonstrated by tuberculin skin testing or an interferon gamma release assay (IGRA): so-called latent TB.
- If these reparative processes fail, primary progressive disease ensues.
- half of this risk occurring in the first 2 years after infection.

# Tuberculosis

## ❖ Types of pulmonary tuberculosis :-

- **Primary tuberculosis;** usually in children, refers to the infection of a previously uninfected (tuberculin-negative) individual.
- May fate to any of the following:-
  - ⇒ **Healing and calcification.**
  - ⇒ **Progressive pulmonary tuberculosis.**
  - ⇒ **Post primary ( secondary tuberculosis).** refers to exogenous (new infection) or most commonly endogenous (reactivation ) of dominant primary infection. Is characterized by localized in apices of one or both upper lobes and the upper segments of lower lobes ( high O<sub>2</sub> concentration ).
  - ⇒ **Miliary tuberculosis.** Blood dissemination of tuberculin bacilli give rise to military, may occur in young adult and old age as a complication of primary tuberculosis.

# Tuberculosis

❖ **Predisposing factors :-** That lower the body resistance.

- **Environmental factors :-** such as malnutrition, poverty, overcrowding, unhygienic condition, alcoholism and heavy smoking.
- **Pathological factors :-** such as DM, steroid, chronic lung disease, lymphoma, and cytotoxic drugs.

# Tuberculosis

## ❖ *Clinical features:*

### ☐ *pulmonary disease*

#### ➤ **Primary pulmonary TB**

- A few patients develop a self-limiting febrile illness but clinical disease occurs only if there is a hypersensitivity reaction or progressive infection.

#### ➤ **Progressive primary disease**

- May appear during the course of the initial illness or after a latent period of weeks or months.



# Tuberculosis

## ❖ Clinical features:

### ☐ pulmonary disease

#### ➤ Post-primary pulmonary TB

- It is most frequently pulmonary and characteristically occurs in the apex of an upper lobe, where the oxygen tension favors survival of the strictly aerobic organism.
- The onset is usually insidious, developing slowly over several weeks.
- Systemic symptoms include fever, night sweats, malaise and loss of appetite and weight, and are accompanied by progressive pulmonary symptoms.
- Very occasionally, this form of TB may present with one of the complications listed below.

# Tuberculosis

## ❖ Clinical features:

### ☐ pulmonary disease

#### ➤ Post-primary pulmonary TB

- Radiological changes include ill-defined opacification in one or both of the upper lobes, and as progression occurs, consolidation, collapse and cavitation develop to varying degrees.
- It is often difficult to distinguish active from quiescent disease on radiological criteria alone but the presence of a miliary pattern or cavitation favors active disease.
- In extensive disease, collapse may be marked and results in significant displacement of the trachea and mediastinum.
- Occasionally, a caseous lymph node may drain into an adjoining bronchus, leading to tuberculous pneumonia.

# Tuberculosis

## ❖ *Clinical features:*

### ☐ *pulmonary disease*

#### ➤ **Miliary TB**

- May present acutely but more frequently is characterized by 2–3 weeks of fever, night sweats, anorexia, weight loss and a dry cough.
- Hepatosplenomegaly may develop and the presence of a headache may indicate coexistent tuberculous meningitis.
- Auscultation of the chest is frequently normal but in more advanced disease widespread crackles are evident.

# Tuberculosis

## ❖ *Clinical features:*

### ☐ *pulmonary disease*

#### ➤ **Miliary TB**

- Fundoscopy may show choroidal tubercles.
- The classical appearances on chest X-ray are of fine 1–2 mm lesions ('millet seed') distributed throughout the lung fields, although occasionally the appearances are coarser.
- Anaemia and leucopenia reflect bone marrow involvement.

# Tuberculosis

## ❖ *Clinical features:*

### ➤ Complications of chronic pulmonary Tuberculosis

#### **Pulmonary**

- Massive haemoptysis
- Cor pulmonale
- Fibrosis/emphysema
- Atypical mycobacterial infection
- Lung/pleural calcification
- Aspergilloma/chronic aspergillosis
- Obstructive airways disease
- Bronchiectasis
- Bronchopleural fistula

#### **Non-pulmonary**

- Empyema necessitans
- Laryngitis
- Enteritis
- Anorectal disease
- Amyloidosis
- Poncet's polyarthrititis

# Tuberculosis

## ❖ *Clinical features:*

- ❑ ***Extrapulmonary disease;*** Accounts for 20% of cases in those who are HIV-negative but is more common in HIV-positive patients.
- **Lymphadenitis**
  - Lymph nodes are the most common extrapulmonary site of disease.
  - Cervical and mediastinal glands are affected most frequently, followed by axillary and inguinal.
  - May represent primary infection, spread from contiguous sites or reactivation.

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Lymphadenitis**

- The nodes are usually painless and initially mobile but become matted together with time.
- Approximately half of cases fail to show any constitutional features.
- The tuberculin test is usually strongly positive.
- During or after treatment, paradoxical enlargement, development of new nodes and suppuration may all occur but without evidence of continued infection.

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Gastrointestinal tuberculosis**

- TB can affect any part of the bowel.
- Upper gastrointestinal tract involvement is rare.
- Ileocecal disease accounts for approximately half of abdominal TB cases.
- Fever, night sweats, anorexia and weight loss are usually prominent and a right iliac fossa mass may be palpable.
- Up to 30% of cases present with an acute abdomen.



# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Gastrointestinal tuberculosis**

- Ultrasound or CT may reveal thickened bowel wall, abdominal lymphadenopathy, mesenteric thickening or ascites.
- Barium enema and small bowel enema reveal narrowing, shortening and distortion of the bowel, with caecal involvement predominating.
- Diagnosis rests on obtaining histology by either colonoscopy or mini-laparotomy.
- The main differential diagnosis is Crohn's disease.
- Tuberculous peritonitis is characterized by abdominal distension, pain and constitutional symptoms.

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Gastrointestinal tuberculosis**

- The ascitic fluid is exudative and cellular, with a predominance of lymphocytes.
- Laparoscopy reveals multiple white 'tubercles' over the peritoneal and omental surfaces.
- Low-grade hepatic dysfunction is common in miliary disease, in which biopsy reveals granulomas.
- Occasionally, patients may be frankly icteric, with a mixed hepatic/cholestatic picture.

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Pericardial disease**

- Disease occurs in two forms: pericardial effusion and constrictive pericarditis.
- Fever and night sweats are rarely prominent and the presentation is usually insidious, with breathlessness and abdominal swelling.
- Coexistent pulmonary disease is very rare, with the exception of pleural effusion.
- Pulsus paradoxus, a raised JVP, hepatomegaly, prominent ascites and peripheral oedema are common to both types.

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Pericardial disease**

- Pericardial effusion is associated with increased pericardial dullness and a globular enlarged heart on chest X-ray, and pericardial calcification occurs in around 25% of cases.
- Constriction is associated with an early third heart sound and, occasionally, atrial fibrillation.
- Diagnosis is based on the clinical, radiological and echocardiographic findings.
- The effusion is frequently blood-stained.
- The addition of glucocorticoids to anti-tuberculosis treatment has been shown to help both forms of pericardial disease.

# Tuberculosis

## ❖ *Clinical features:*

❑ *Extrapulmonary disease;*

### ➤ **Central nervous system disease:-**

- **Meningeal disease represents the most important form of central nervous system TB.**
- **Unrecognised and untreated, it is rapidly fatal.**
- **Even when appropriate treatment is prescribed, mortality rates of 30% have been reported, while survivors may be left with neurological sequelae.**

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Bone and joint disease**

- The spine is the most common site for bony TB (Pott's disease).
- Usually presents with chronic back pain and typically involves the lower thoracic and lumbar spine.
- The infection starts as a discitis and then spreads along the spinal ligaments to involve the adjacent anterior vertebral bodies, causing angulation of the vertebrae with subsequent kyphosis.

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Bone and joint disease**

- Paravertebral and psoas abscess formation is common and the disease may present with a large (cold) abscess in the inguinal region.
- CT or MRI is valuable in gauging the extent of disease, the amount of cord compression, and the site for needle biopsy or open exploration, if required.
- The major differential diagnosis is malignancy.
- Important complications include spinal instability or cord compression.

# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Bone and joint disease**

- TB can affect any joint but most frequently involves the hip or knee.
- Presentation is usually insidious, with pain and swelling; fever and night sweats are uncommon.
- Radiological changes are often non-specific but, as disease progresses, reduction in joint space and erosions appear.
- Poncet's arthropathy is an immunologically mediated polyarthrititis that usually resolves within 2 months of starting treatment.



# Tuberculosis

## ❖ *Clinical features:*

### ❑ *Extrapulmonary disease;*

#### ➤ **Genitourinary disease**

- Fever and night sweats are rare with renal tract TB and patients are often only mildly symptomatic for many years.
- Haematuria, frequency and dysuria are often present, with sterile pyuria found on urine microscopy and culture.
- In women, infertility from endometritis, or pelvic pain and swelling from salpingitis or a tubo-ovarian abscess occurs occasionally.
- In men, genitourinary TB may present as epididymitis or prostatitis.

# Tuberculosis

## ❖ *Investigations*

- The presence of an otherwise unexplained cough for more than 2–3 weeks, particularly in regions where TB is prevalent, or typical chest X-ray or CT changes should prompt further investigation.
- **X ray ( PA view ) :-**
  - ⇒ In early stage ill-defined opacities usually in one of the upper lobe.
  - ⇒ In advanced stage opacities are larger, widespread and may be bilateral.
  - ⇒ Cavitation.
  - ⇒ Displaced trachea and heart shadow especially in complication.

# Tuberculosis

## ❖ *Investigations*

- **Sputum for AFB :-**

- **Direct microscopy of a sputum smear remains the most important first step.**
- **Microscopic ex at least 3 specimens of sputum, prefer collected earl in the morning.**
- **At least two sputum samples (including at least one obtained in the early morning) from a spontaneously produced deep cough should be obtained.**
- **Induced sputum may be used in those unable to expectorate.**

# Tuberculosis

## ❖ *Investigations*

- **Sputum for AFB :-**

- In selected cases, bronchoscopy and lavage or aspiration of a lymph node may be used.
- A positive smear is sufficient for the presumptive diagnosis of TB but definitive diagnosis requires culture.
- The probability of detecting acid-fast bacilli is proportional to the bacillary burden in the sputum.

# Tuberculosis

## ❖ *Investigations*

- **Culture**

- Smear-negative sputum should also be cultured, as only 10–100 viable organisms are required for sputum to be culture-positive.
- A diagnosis of smear-negative TB may be made in advance of culture if the chest X-ray appearances are typical of TB.
- The slow growth of MTB on solid (typically between 4 and 6 weeks) and automated and semi-automated liquid (typically around 2 weeks) culture media has prompted the development of rapid NAATs

# Tuberculosis

## ❖ *Investigations*

- **Monteux tuberculin skin test :-**

- ⇒ **Monteux test with purified protein derivatives PPD injectable to skin, the transverse width mm of the induration should be recorded after 48 72 hrs.**
- ⇒ **MT used to screen active disease and latent infection, but does not distinguish between them.**
- ⇒ **A negative test does not rule out the diagnosis of TB, but a positive test may be helpful for diagnosis.**
- ⇒ **Test is positive if the induration is 10 or more in diameter.**

# Tuberculosis

## ❖ *Investigations*

- **Monteux tuberculin skin test :-**

- ⇒ **MT also considered positive if more than 5 for patient with HIV, close contact with active TB or immunocompromised.**
- ⇒ **False positive in infection with non-tuberculosis mycobacterium.**
- ⇒ **False negative may be due to weak immunity, immunodeficiency state, and improper testing technique.**

# Tuberculosis

## ❖ *Investigations*

- **NAAT**
  - (a DNA detection-based NAAT) has the capacity to detect MTB in less than 2 hours.
  - It is specific to MTB, it is not sufficiently sensitive to have replaced culture.
- **Interferon gamma release assay ( IGRA ):-** with antigen specific for mycobacterium tuberculosis also used to screen latent TB.
- **CBC, ESR, CRP.**
- **Heaf test and tine test :-** this simple test used for screening.



# Tuberculosis

## ❖ *Investigations*

- **Needle biopsy** :- from pleural, lymph nodes, and solid lesion within the lung and culture.
- **Pleural fluid aspiration** :- predominant lymphocyte, and culture is positive in less than 20%.
- **PCR** :- it is common and rapid test for detection of mycobacterial DNA in sputum and other fluids within 48 hrs.

# Tuberculosis

## ❖ *Investigations*

- The diagnosis of extrapulmonary TB can be more challenging.
- Generally fewer organisms (particularly in meningeal or pleural fluid).
- Culture, histopathological examination of tissue and/or NAAT may be required.
- Adenosine deaminase in pleural, pericardial, cerebrospinal and ascitic fluid, and so may assist in confirming suspected TB.
- In the presence of HIV, examination of sputum may still be useful, as subclinical pulmonary disease is common.
- Lateral flow urinary lipoarabinomannan assay (LF-LAM) may be useful in the severely ill patient with a CD4 count of 100 cells/ $\mu$ L or less.

# Tuberculosis

## ❖ *Management*

- **General measures :-**

- ⇒ Isolate patients with possible TB in private room with negative pressure.
- ⇒ Have medical staff wear high efficiency disposable masks.
- ⇒ Continue isolation until sputum smears are negative for 3 consecutive times ( usually after 2 4 wks. of treatment ).

# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ The treatment of TB is based on the principle of an initial intensive phase to reduce the bacterial population rapidly, followed by a continuation phase to destroy any remaining bacteria.
- ✓ Standard treatment involves 6 months' treatment with isoniazid and rifampicin, supplemented in the first 2 months with pyrazinamide and ethambutol.
- ✓ Fixed-dose tablets combining two or three drugs are preferred.

# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ **Treatment should be commenced immediately in any patient who is smear-positive, and in those who are smear-negative but with typical chest X-ray changes and no response to standard antibiotics.**
- ✓ **12 months of therapy is recommended for meningeal TB, including involvement of the spinal cord in cases of spinal TB; in these cases, ethambutol may be replaced by streptomycin.**
- ✓ **Pyridoxine should be prescribed in pregnant women and malnourished patients to reduce the risk of peripheral neuropathy with isoniazid.**

# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ **Where drug resistance is not anticipated, patients can be assumed to be non-infectious after 2 weeks of appropriate therapy.**
- ✓ **Most patients can be treated at home.**
- ✓ **Six months of therapy is appropriate for all patients with new-onset pulmonary TB and most cases of extrapulmonary TB.**

# Tuberculosis

## ❖ *Management*

### ➤ Chemotherapy

- ✓ Admission to a hospital unit with appropriate isolation facilities should be considered where;
  - Uncertainty about the diagnosis.
  - Intolerance of medication.
  - Questionable treatment adherence.
  - Adverse social conditions.
  - A significant risk of multidrug-resistant TB (culture positive after 2 months on treatment, or contact with known multidrug-resistant TB).

# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ Patients treated with rifampicin should be advised that their urine, tears and other secretions will develop a bright, orange/ red coloration.
- ✓ Women taking the oral contraceptive pill must be warned that its efficacy will be reduced and alternative contraception may be necessary.
- ✓ Ethambutol and streptomycin should be used with caution in renal impairment, with appropriate dose reduction and monitoring of drug levels.
- ✓ Adverse drug reactions occur in about 10% of patients but are significantly more common with HIV co-infection.



# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ **Baseline liver function and regular monitoring are important for patients treated with standard therapy.**
- ✓ **Rifampicin may cause asymptomatic hyperbilirubinemia but, along with isoniazid and pyrazinamide, may also cause hepatitis.**
- ✓ **Mild asymptomatic increases in transaminases are common but significant hepatotoxicity only occurs in 2–5%.**
- ✓ **It is appropriate to stop treatment and allow any symptoms to subside and the liver function tests to recover before commencing a stepwise re-introduction of the individual drugs.**

# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ Less hepatotoxic regimens may be considered, including streptomycin, ethambutol and fluoroquinolones.
- Streptomycin and pyrazinamide contraindication in pregnancy.
- All anti TB are safe for lactating mother and is not harmful to nursing neonate.
- In renal impairment avoid ethambutol, and aminoglycosides.
- In severe liver disease best used she thereby.

# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ **Glucocorticoids are currently recommended in pericardial or meningeal disease.**
- ✓ **May benefit in TB of the ureter, pleural effusions and extensive pulmonary disease.**
- ✓ **Can suppress hypersensitivity drug reactions.**
- ✓ **Surgery should be considered in cases complicated by massive hemoptysis, loculated empyema, constrictive pericarditis, lymph node suppuration, and spinal disease with cord compression, but usually only after a full course of anti-tuberculosis treatment.**

# Tuberculosis

## ❖ *Management*

### ➤ **Chemotherapy**

- ✓ **The effectiveness of therapy for pulmonary TB is assessed by further sputum smear at 2 months and at 5 months.**
- ✓ **Treatment failure is defined as a positive sputum smear or culture at 5 months or any patient with a multidrug-resistant strain, regardless of whether they are smear-positive or negative.**
- ✓ **Extrapulmonary TB must be assessed clinically or radiographically, as appropriate.**

# Tuberculosis

## ❖ *Management*

### ➤ Monitoring response to treatment :-

- Clinical assessment.
- Sputum for AFB smear monthly until become negative.
  - ⇒ By the end of 3 rd. months of treatment all patient should be smear negative.
  - ⇒ If smear remain positive at or beyond 3 months, treatment failure or drug resistance should be suspected.
- Chest x ray is performed initially and at the end of treatment for comparison.
- A base line LFT because hepatitis is a major complication of anti TB.

# Tuberculosis

## ❖ *Management*

### ➤ **Treatment failure:-**

- Said when the patients sputum culture remain positive after 3 months, or AFB smear remain positive after 5 months.
- Drug resistance usually develops when patient uses single drug thereby or irregular tokening properly prescribed thereby.
- In this cases added at least 2 or 3 drugs that have never been used before.

# Tuberculosis

## ❖ *Management*

### ➤ **Directly observed therapy**

- ✓ **Poor adherence to therapy is a major factor in prolonged illness, risk of relapse, and the emergence of drug resistance.**
- ✓ **Directly observed therapy (DOT) involves the supervised administration of therapy 3 times weekly to improve adherence.**
- ✓ **DOT has become an important control strategy in resource-poor nations.**
- ✓ **It is currently recommended for patients thought unlikely to be adherent to therapy: homeless people and drifters, alcohol or drug users, patients with serious mental illness and those with a history of non-adherence.**

# Tuberculosis

## ❖ *Management*

### ➤ *Control and prevention*

- ✓ TB is preventable, particularly so in those with latent TB.
- ✓ Supporting the development of laboratory and health-care services to improve detection and treatment of active and latent TB is an important component of this goal.
- ✓ The majority of individuals exposed to MTB harbour the bacteria, which remain dormant.
- ✓ They do not develop any signs of active disease and are non-infectious.



# Tuberculosis

## ❖ *Management*

### ➤ *Control and prevention*

- ✓ They are however, at risk of developing active TB disease and becoming infectious.
- ✓ The lifetime risk of TB disease for a person with documented latent TB infection is estimated at 5–15%, with the majority of cases occurring within the first 5 years after initial infection.
- ✓ Latent TB may be identified by the presence of immune responses to *M. tuberculosis* antigens.
- ✓ Contact tracing is a legal requirement in many countries.

# Tuberculosis

## ❖ *Management*

### ➤ *Control and prevention*

- ✓ Close contacts who should receive BCG vaccination or chemotherapy.
- ✓ Approximately 10–20% of close contacts of patients with smear-positive pulmonary TB and 2–5% of those with smear-negative, culture-positive disease have evidence of TB infection.
- ✓ Cases are commonly identified using the tuberculin skin test (TST) or an IGRA. An otherwise asymptomatic contact who tests positive but has a normal chest X-ray may be treated with chemoprophylaxis to prevent infection from progressing to clinical disease.
- ✓ Chemoprophylaxis should be offered to adults up to the age of 65.

# Tuberculosis

## ❖ *Management*

### ➤ *Control and prevention*

- ✓ It should also be considered for HIV-infected close contacts of a patient with smear-positive disease.
- ✓ A course of rifampicin and isoniazid for 3 months or isoniazid for 6 months is effective.
- ✓ Tuberculin skin testing may be associated with false-positive reactions in those who have had a BCG vaccination and in areas where exposure to non-tuberculous mycobacteria is high.
- ✓ The skin tests may also be falsely negative in the setting of immunosuppression or overwhelming TB infection.

# Tuberculosis

## ❖ Management

### ➤ Control and prevention

- ✓ IGRAs detect the release of interferon-gamma (IFN- $\gamma$ ) from sensitized T cells in response to antigens, such as early secretory antigenic target (ESAT)-6 or culture filtrate protein (CFP)-10, which are encoded by genes specific to *Mycobacterium tuberculosis* and are not shared with BCG or opportunistic mycobacteria.
- ✓ IGRAs are more specific than skin testing as they require a single blood test rather than two clinic visits.
- ✓ A dual strategy of TST followed by IGRA is recommended.
- ✓ TST remains the first choice in children, while IGRA represents the first choice for individuals with HIV

# Tuberculosis

## ❖ Management

### ➤ Control and prevention

#### ❑ Vaccines

- ✓ BCG (the Calmette–Guérin bacillus), a live attenuated vaccine derived from *M. bovis*, is the most established TB vaccine.
- ✓ It is administered by intradermal injection and is highly immunogenic.
- ✓ BCG appears to be effective in preventing disseminated disease, including tuberculous meningitis, in children, but its efficacy in adults is inconsistent and new vaccines are urgently needed.
- ✓ Current vaccination policies usually target children and other high-risk individuals.
- ✓ BCG is very safe, with the occasional complication of local abscess formation.
- ✓ Not be administered to those who are immunocompromised (e.g. by HIV) or pregnant.

# Tuberculosis

## ❖ *Management*

### ➤ TB and HIV/AIDS

- ✓ The close links between HIV and TB, with the promotion of programs that link detection and treatment of TB with detection and treatment of HIV.
- ✓ It is recommended that all patients with TB should be tested for HIV infection.
- ✓ Mortality is high and TB is a leading cause of death in HIV patients.

# Tuberculosis

## ❖ *Management*

### ➤ Drug-resistant TB

- ❑ **Drug-resistant TB** is defined by the presence of resistance to any first-line agent.
- ❑ **Multidrug-resistant tuberculosis (MDR-TB)** is defined by resistance to at least rifampicin and isoniazid, with or without other drug resistance.
- ❑ **Extensively drug-resistant tuberculosis (XDR-TB)** is defined as resistance to at least rifampicin and isoniazid, in addition to any quinolone and at least one injectable second-line agent.

# Tuberculosis

## ❖ *Management*

### ➤ Drug-resistant TB

- ✓ More common in individuals with a prior history of TB, particularly if treatment has been inadequate, and those with HIV infection.
- ✓ Diagnosis is challenging, especially in resource-poor settings, and although cure may be possible, it requires prolonged treatment with less effective, more toxic and more expensive therapies.
- ✓ When MDR-TB is suspected, administration at least 5 drugs for intensive phase of Treatment and at least 4 drugs for the continues phase.
- ✓ The mortality rate from MDR-TB is high and that from XDR-TB higher still.



# Tuberculosis

## ❖ *Management*

### ➤ Drug-resistant TB

- ✓ The factors contributing to the emergence of drug-resistant TB.
  - Drug shortages.
  - Poor-quality drugs.
  - Lack of appropriate supervision.
  - Transmission of drug-resistant strains.
  - Prior anti-tuberculosis treatment.
  - Treatment failure (smear-positive at 5 months).

# Tuberculosis

## ❖ *Prognosis*

- ✓ Following successful completion of chemotherapy, cure should be anticipated in the majority of patients.
- ✓ There is a small ( $< 5\%$ ) and unavoidable risk of relapse.
- ✓ Most relapses occur within 5 months and usually have the same drug susceptibility.
- ✓ In the absence of treatment, a patient with smear-positive TB will remain infectious for an average of 2 years; in 1 year, 25% of untreated cases will die.

# Tuberculosis

## ❖ *Prognosis*

- ✓ Death is more likely in those who are smear-positive and those who smoke.
- ✓ A few patients die unexpectedly soon after commencing therapy and it is possible that some have subclinical hypoadrenalism that is unmasked by a rifampicin-induced increase in glucocorticoid metabolism.
- ✓ HIV-positive patients have higher mortality rates and a modestly increased risk of relapse.



# Thank you